

Serial No. 09/928,262

IN THE CLAIMS:

1. (Currently amended) A method of delivering a nucleic acid of interest to a primary human chondrocyte, comprising:
providing a recombinant chimeric adenovirus having a tropism for primary human chondrocytes,
said recombinant chimeric adenovirus comprising:
a nucleic acid of interest operatively linked to a promoter, wherein said nucleic acid of interest encodes at least one amino acid sequence that inhibits cartilage disease progression, at least one amino acid sequence that counteracts the loss of cartilage, or a combination thereof;
a deletion in a gene encoding a fiber protein; and
a nucleic acid replacing the deletion in the gene of the fiber protein, said nucleic acid encoding at least a part of a fiber protein of a B-type adenovirus;
wherein said at least a part of the fiber protein of the B-type adenovirus has a tropism for primary human chondrocytes; and
infecting a primary human chondrocyte in vitro with said recombinant chimeric adenovirus, such that said nucleic acid of interest is delivered to said primary human chondrocyte.
- 2-4. Canceled.
5. (Previously presented) The method of claim 1, wherein said B-type adenovirus is adenovirus type 35.
6. (Previously presented) The method of claim 1, wherein said recombinant chimeric adenovirus comprises an adenovirus 5 nucleic acid sequence.
7. (Previously presented) The method of claim 5, wherein said recombinant chimeric adenovirus comprises an adenovirus 5 genome.

Serial No. 09/928,262

8. (Previously presented) The method of claim 1, wherein said recombinant chimeric adenovirus comprises at least one deletion in the E3 region where the nucleic acid of interest is inserted or can be inserted.

9-23. Canceled

24. (Currently amended) ~~Chondrocytes~~ In vitro chondrocytes provided with an additional nucleic acid encoding:
at least one amino acid sequence that inhibits cartilage disease progression;
at least one amino acid sequence that counteracts the loss of cartilage; or
a combination thereof;
said additional nucleic acid provided by a gene delivery vehicle comprising a recombinant chimeric adenovirus having a tropism for chondrocytes;
said recombinant chimeric adenovirus comprising:
a deletion in a gene encoding a fiber protein; and
a nucleic acid replacing the deletion in the gene encoding the fiber protein, said nucleic acid encoding at least a part of a fiber protein of a B-type adenovirus;
wherein said at least a part of the fiber protein of the B-type adenovirus has a tropism for primary human chondrocytes.

25. (Currently amended) The in vitro chondrocytes of claim 24, wherein said additional nucleic acid encodes at least one member of the family of bone morphogenesis proteins.

26. Canceled.

Serial No. 09/928,262

27. (Currently amended) A method of inhibiting cartilage disease progression comprising:
preparing a recombinant chimeric adenovirus having a tropism for primary human chondrocytes,
said recombinant chimeric adenovirus including:
a nucleic acid encoding a protein useful in inhibiting cartilage disease progression operatively
linked to a promoter;
a deletion in a gene encoding a fiber protein; and
a nucleic acid replacing the deletion in the gene encoding the fiber protein, said nucleic acid
encoding at least a part of a fiber protein of a B-type adenovirus;
wherein said at least a part of the fiber protein of the B-type adenovirus has a tropism for primary
human chondrocytes; and
infecting a primary human chondrocyte in vitro with said recombinant chimeric adenovirus, such
that said nucleic acid of interest encoding the protein useful in inhibiting cartilage disease
progression is expressed in said primary human chondrocyte, inhibiting cartilage disease
progression.

28. (Currently amended) A method of repairing cartilage comprising:
preparing a recombinant chimeric adenovirus having a tropism for primary human chondrocytes,
said recombinant chimeric adenovirus including:
a nucleic acid encoding a protein useful in repairing cartilage operatively linked to a promoter;
a deletion in a gene encoding a fiber protein; and
a nucleic acid replacing the deletion in the gene encoding the fiber protein, said nucleic acid
encoding at least a part of a fiber protein of a B-type adenovirus;
wherein said at least a part of the fiber protein of the B-type adenovirus has a tropism for primary
human chondrocytes; and
infecting a primary human chondrocyte in vitro with said recombinant adenovirus, such that said
nucleic acid encoding the protein useful in repairing cartilage is expressed in said primary
human chondrocyte, effecting the cartilage repair.

Serial No. 09/928,262

29. (New) A method of delivering a nucleic acid of interest to a primary human chondrocyte, comprising:
providing a recombinant chimeric adenovirus having a tropism for primary human chondrocytes,
said chimeric recombinant adenovirus comprising:
a nucleic acid of interest operatively linked to a promoter, wherein said nucleic acid of interest
encodes at least one member of the family of bone morphogenesis proteins;
a deletion in a gene encoding a fiber protein; and
a nucleic acid replacing the deletion in the gene of the fiber protein, said nucleic acid encoding at
least a part of the fiber protein of a B-type adenovirus, wherein said at least a part of the
fiber protein of the B-type adenovirus has a tropism for primary human chondrocytes;
and
infecting a primary human chondrocyte in vitro with said recombinant chimeric adenovirus, such
that said nucleic acid of interest is delivered to said primary human chondrocyte.

30. (New) In a method of infecting a cell with an adenoviral vector having a
recombinant fiber protein comprising a part of a fiber protein originating from an Ad5 fiber
protein and an adenoviral genome, the improvement comprising:
wherein a part of the recombinant fiber protein originates from a B-type adenovirus fiber protein,
thus providing the recombinant fiber protein with a tissue tropism for primary human
chondrocytes; and
the adenoviral genome further comprises a nucleic acid of interest operatively linked to a
promoter, wherein the nucleic acid of interest encodes at least one amino acid sequence
that inhibits cartilage disease progression, at least one amino acid sequence that
counteracts the loss of cartilage, or a combination thereof.